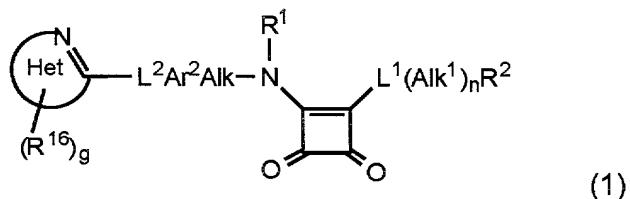


CLAIMS

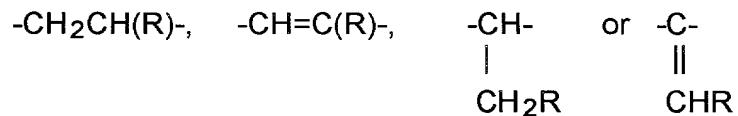
1. A compound of formula (1):

5



wherein

Het is a bicyclic fused ring heteroaromatic group;
 10 g is zero or the integer 1, 2, 3 or 4;
 Each R¹⁶, which may be the same or different is an atom or group -L³(Alk²)_tL⁴(R⁴)_u in which L³ and L⁴, which may be the same or different, is each a covalent bond or a linker atom or group, t is zero or the integer 1, u is an integer 1, 2 or 3, Alk² is an aliphatic or heteroaliphatic chain and R⁴ is a hydrogen or halogen atom or a group selected from optionally substituted C₁₋₆alkyl or C₃₋₈ cycloalkyl, -OR⁵ [where R⁵ is a hydrogen atom, an optionally substituted C₁₋₆alkyl or C₃₋₈ cycloalkyl group], -SR⁵, -NR⁵R⁶ [where R⁶ is as just defined for R⁵ and may be the same or different], -NO₂, -CN, -CO₂R⁵, -SO₃H, -SOR⁵, -SO₂R⁵, -SO₃R⁵, -OCO₂R⁵, -CONR⁵R⁶, -OCONR⁵R⁶, -CSNR⁵R⁶, -COR⁵, -OCOR⁵, -N(R⁵)COR⁶, -N(R⁵)CSR⁶, -SO₂N(R⁵)(R⁶), -N(R⁵)SO₂R⁶, N(R⁵)CON(R⁶)(R⁷) [where R⁷ is a hydrogen atom, an optionally substituted C₁₋₆alkyl or C₃₋₈cycloalkyl group], -N(R⁵)CSN(R⁶)(R⁷) or -N(R⁵)SO₂N(R⁶)(R⁷), provided that when t is zero and each of L³ and L⁴ is a covalent bond then u is the integer 1 and R⁴ is other than a hydrogen atom;
 15 L² is a covalent bond or an atom or group -O-, -S-, -C(O)-, -C(S)-, -S(O)-, -S(O)₂, -N(R⁸)- [where R⁸ is a hydrogen atom or an optionally substituted C₁₋₆alkyl group] or -C(R⁸)(R^{8a})- [where R^{8a} is an atom or group as defined for R⁸ and may be the same or different];
 20 Ar² is an optionally substituted aromatic or heteroaromatic group;
 25 Alk is a chain
 30



in which R is a carboxylic acid (-CO₂H) or a derivative or biostere thereof;

R¹ is a hydrogen atom or a C₁-6alkyl group;

L^1 is a covalent bond or a linker atom or group;

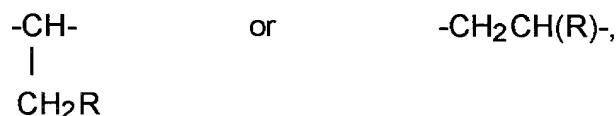
Alk¹ is an optionally substituted aliphatic chain;

n is zero or the integer 1;

10 R² is a hydrogen atom or an optionally substituted heteroaliphatic, cycloaliphatic, heterocycloaliphatic, polycycloaliphatic, heteropolycyclo-aliphatic, aromatic or heteroaromatic group; provided that Het is not a 2,6-naphthyridin-1-yl, isoquinolin-1-yl, 2,7-naphthyridin-1-yl or quinazolin-4-yl group;

15 and the salts, solvates, hydrates and N-oxides thereof.

2. A compound according to Claim 1 in which Alk is a chain



3. A compound according to Claim 1 in which R is a carboxylic acid (-CO₂H) group.

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4. A compound according to Claim 1 in which R is an esterified carboxyl group of formula -CO₂Alk⁷.

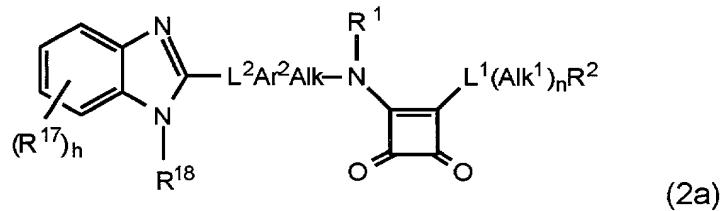
5. A compound according to Claim 1 in which R¹ is a hydrogen atom.

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6. A compound according to Claim 1 in which Ar² is an optionally substituted phenylene group.

35 7. A compound according to Claim 1 in which L¹ is a -N(R⁸)- group where R⁸ is a hydrogen atom or an optionally substituted C₁₋₆alkyl group.

8. A compound according to Claim 7 in which R⁸ is a methyl, ethyl or n-propyl group.
9. A compound according to Claim 1 in which L¹ is a covalent bond.
- 5 10. A compound according to Claim 1 in which n is the integer 1, Alk¹ is an optionally substituted straight or branched C₁-6alkylene chain and R² is a hydrogen atom.
- 10 11. A compound according to Claim 10 in which Alk¹ is a -CH₂-, -CH₂CH₂-, -CH₂CH₂CH₂-, -CH(CH₃)CH₂- or -C(CH₃)₂CH₂- chain.
12. A compound according to Claim 1 in which L¹ is a covalent bond, n is zero and R² is an optionally substituted C₅-7heterocycloaliphatic group.
- 15 13. A compound according to Claim 12 in which R² is an optionally substituted piperidinyl, homopiperidinyl, heptamethyleneiminy, pyrrolidinyl, piperazinyl, homopiperazinyl, morpholinyl or thiomorpholinyl group.
- 20 14. A compound according to Claim 1 in which L² is an -O- atom or -N(R⁸)- group in which R⁸ is a hydrogen atom or an optionally substituted C₁-6alkyl group.
- 25 15. A compound according to Claim 1 of formula (2a):



- 30 wherein:
 R¹⁷ is an atom or group R¹⁶ as previously defined;
 g is the integer 1, 2, 3 or 4;

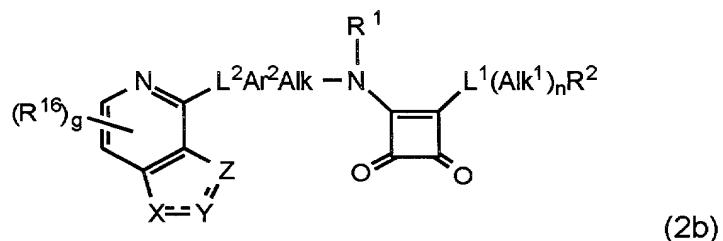
h is zero or the integer 1, 2 or 3;

R^{18} is a hydrogen atom or an atom or group R^{16} as previously defined;

and the salts, solvates, hydrates and N-oxides thereof.

5

16. A compound according to Claim 1 of formula (2b):



10 wherein:

X, Y and Z is each independently selected from a nitrogen, oxygen or sulphur atom or CH group;

the broken line (--) represents saturation or unsaturation;

and the salts, solvates, hydrates and N-oxides thereof.

15

17. A compound according to Claim 16 in which X is an O or S atom, Y and Z are each a group CH, a single bond joins X and Y and a double bond joins Y and Z.

20 18. A compound according to Claim 16 in which Z is an O or S atom, X and Y is each a CH group, a single bond joins Y and Z and a double bond joins X and Y.

19. A compound which is:

25 S-2-{{2-Dipropylamino)-3,4-dioxo-1-cyclobutenyl]amino}-3-{4-[(1-methylbenzimidazol-2-yl)amino]phenyl}propanoic acid;

S-2-{{2-Dipropylamino)-3,4-dioxo-1-cyclobutenyl]amino}-3-{4-[(1-methylbenzimidazol-2-yl)amino]phenyl}propanoic acid;

S-2-{{2-(2-Methylpiperidin-1-yl)-3,4-dioxo-1-cyclobutenyl]amino}-3-{4-

30 [(1-methylbenzimidazol-2-yl)amino]phenyl}propanoic acid;

(S)-3-[4-(Thiophen[2,3-d]pyrimidin-4-ylamino)phenyl]2-(2-diethylamino-3,4-dioxocyclobut-1-enylamino)propanoic acid; and the salts, solvates, hydrates, N-oxides and carboxylic acid esters, particularly the methyl, ethyl, propyl and i-propyl esters thereof.

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20. A pharmaceutical composition comprising a compound according to Claim 1 together with one or more pharmaceutically acceptable carriers, excipients or diluents.
- 10 21. A compound for the prophylaxis or treatment of a disease or disorder in a mammal in which the extravasation of leukocytes plays a role, comprising administering to a mammal suffering from such a disease or disorder a therapeutically effective amount of a compound according to Claim 1.
- 15 22. A method according to Claim 21 wherein said disease or disorder is selected from the group consisting of inflammatory arthritis, multiple sclerosis, allograft rejection, diabetes, inflammatory dermatoses, asthma and inflammatory bowel disease.
- 20 23. A method according to Claim 22 wherein said inflammatory arthritis is selected from the group consisting of rheumatoid arthritis, vasculitis and polydermatomyositis.
- 25 24. A method according to Claim 22 wherein said inflammatory dermatoses are selected from the group consisting of prosiasis and dermatitis.
- 30 25. A method of inhibiting, in a mammal, the binding of $\alpha 4$ integrins to the ligands thereof, comprising administering to the mammal an effecting amount of a compound according to Claim 1.
26. A method according to Claim 25 wherein the $\alpha 4$ integrins are selected from the group consisting of $\alpha 4\beta 1$ and $\alpha 4\beta 7$ integrins.

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